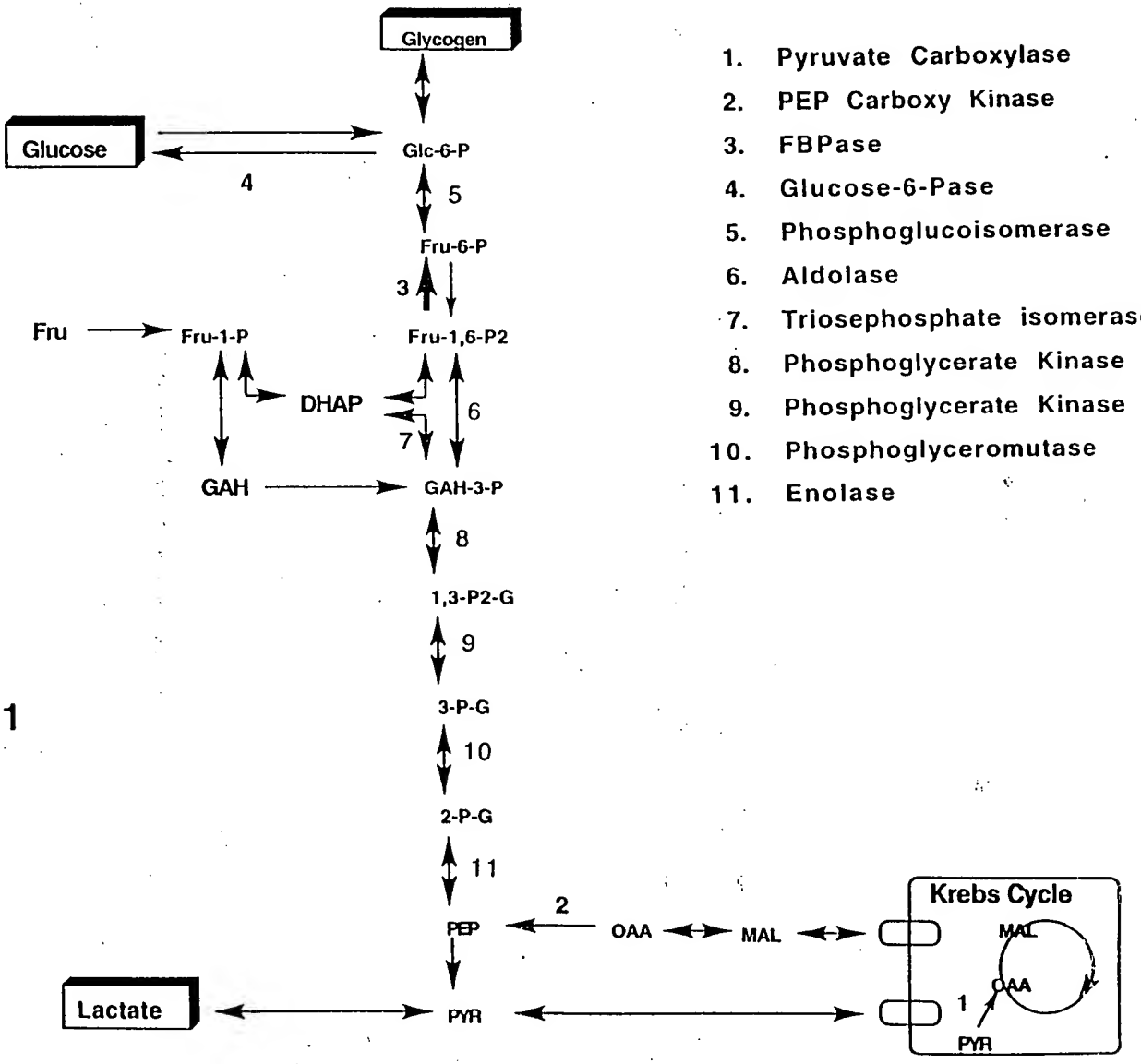

1. Pyruvate Carboxylase
2. PEP Carboxy Kinase
3. FBPase
4. Glucose-6-Pase
5. Phosphoglucoisomerase
6. Aldolase
7. Triosephosphate isomerase
8. Phosphoglycerate Kinase
9. Phosphoglycerate Kinase
10. Phosphoglyceromutase
11. Enolase

FIGURE 1

#4

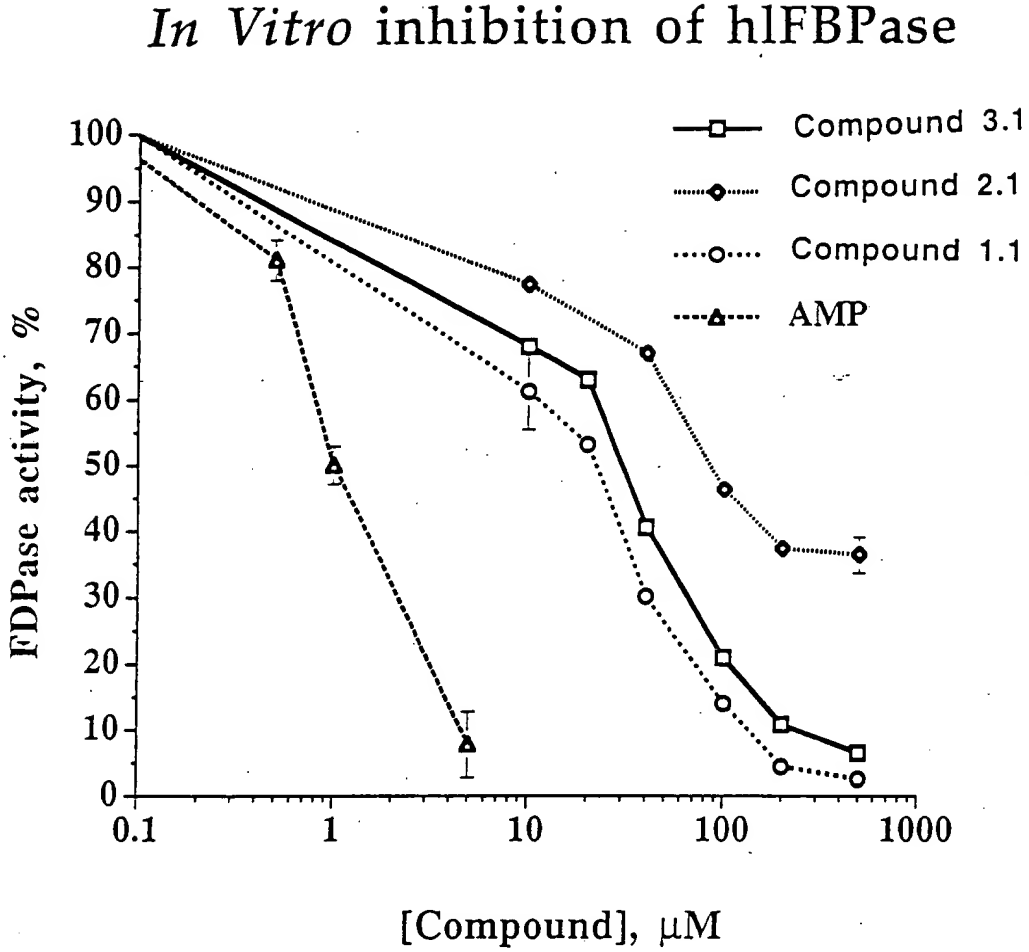
In Vitro inhibition of hlFBPase
Compound 3.1
Compound 2.1
Compound 1.1
AMP
FDPase activity, %
100
90
80
70
60
50
40
30
20
10
0
0.1
1
10
100
1000
[Compound], μM

FIGURE 2

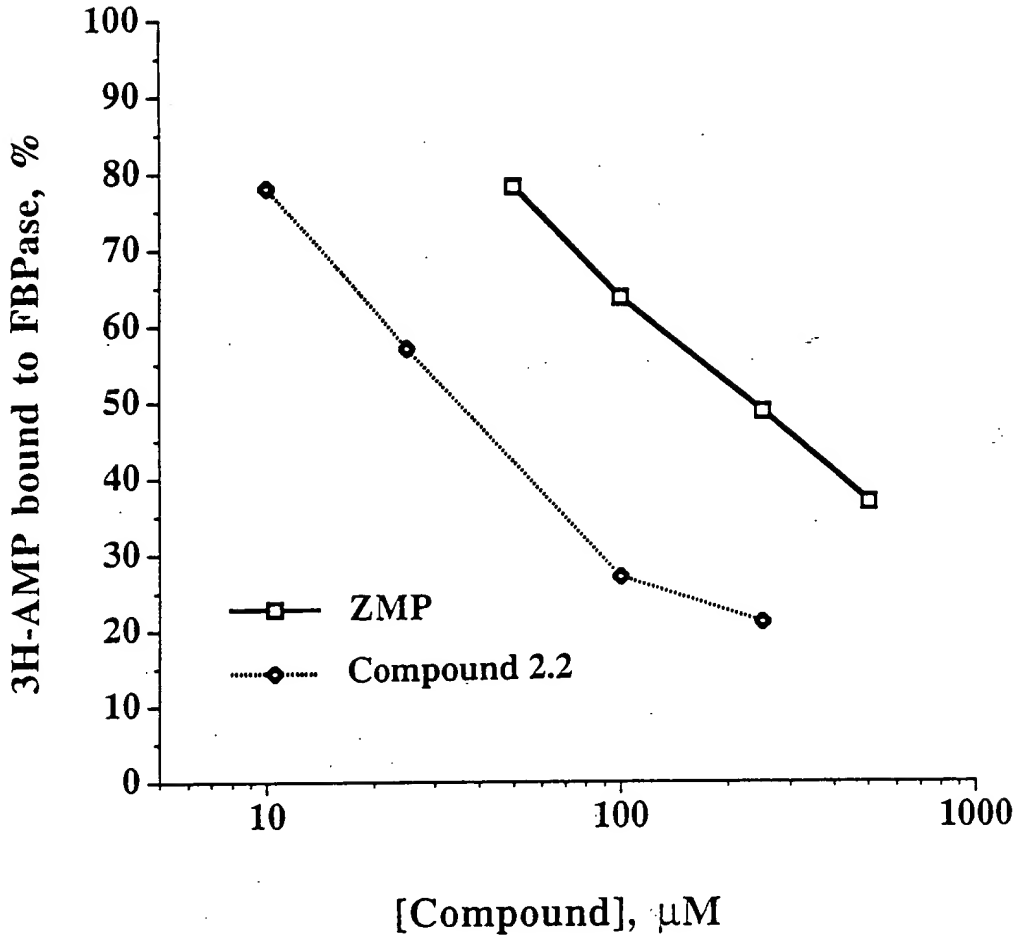
Displacement of AMP from hlFBPase
3H-AMP bound to FBPase, %
100
90
80
70
60
50
40
30
20
10
0
10
100
1000
ZMP
Compound 2.2
[Compound], μM

FIGURE 3

Effect of Compound 2.7 on Gluconeogenesis from Dihydroxyacetone in Rat Hepatocytes

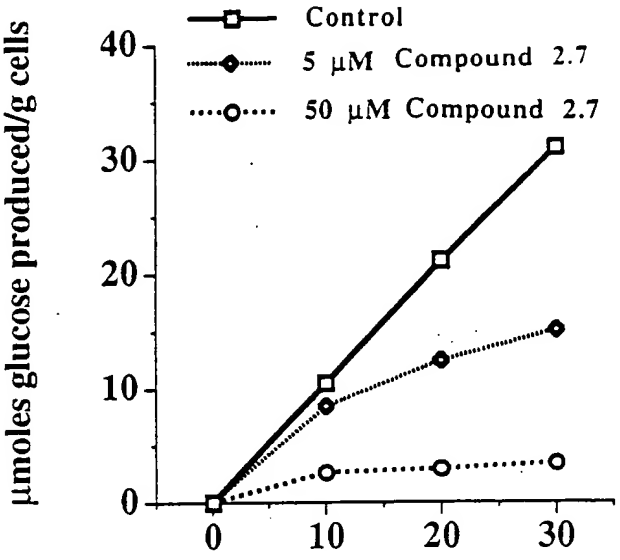

FIGURE 4A

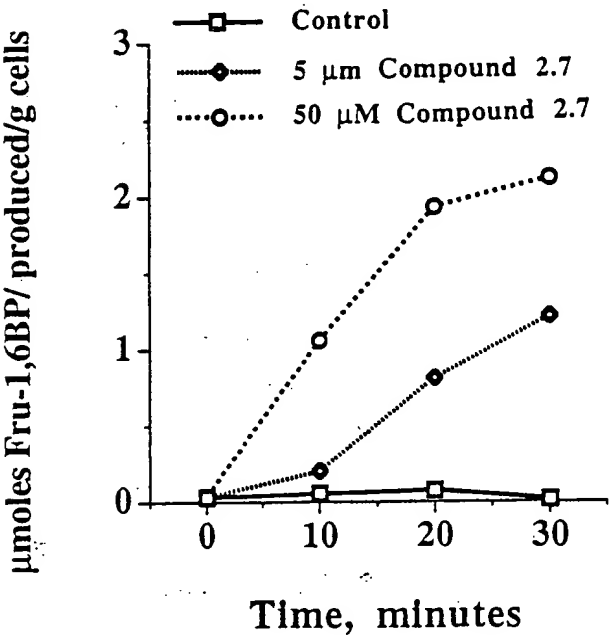

FIGURE 4B

Inhibition of Glucose Production From Lactate Pyruvate (Rat Hepatocytes)

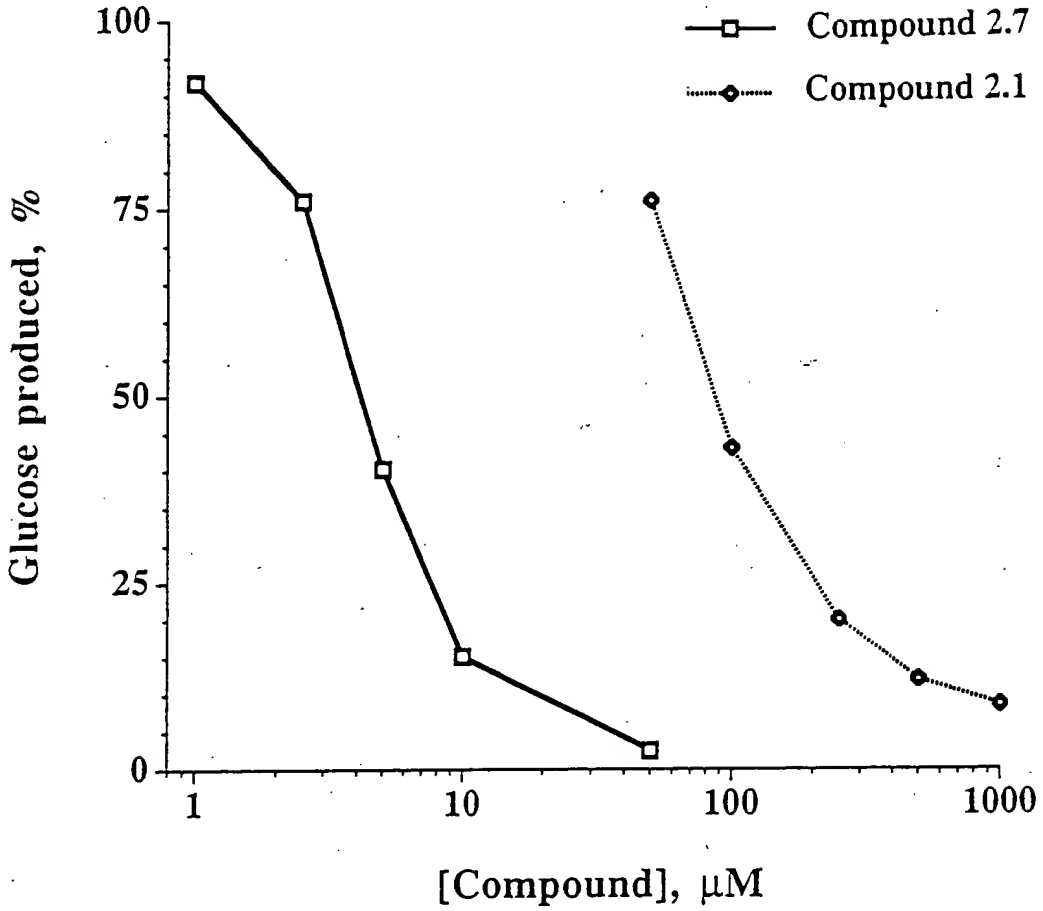

FIGURE 5

Compound 16.4 in 18h-Fasted, Normal Rats (i.p.)

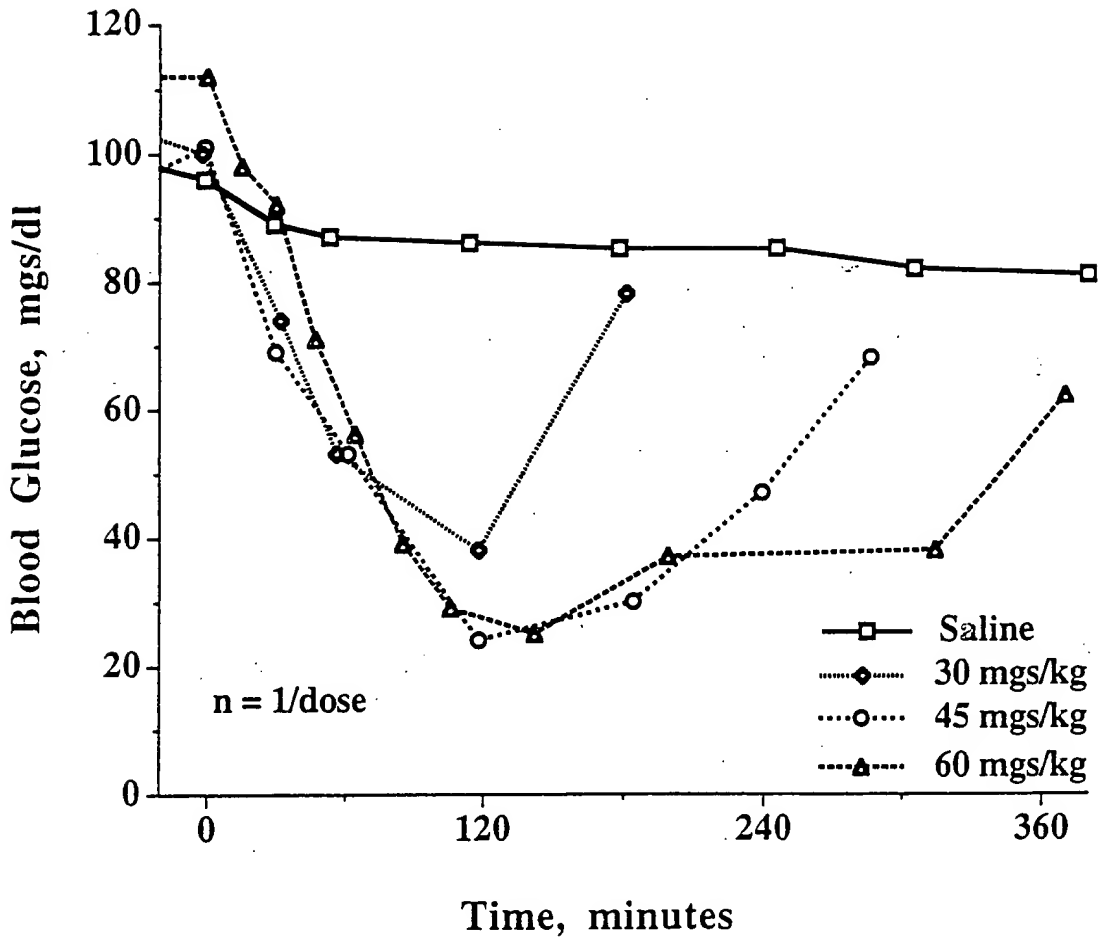

FIGURE 6

Compound 2.7 in 18-hour fasted rats
(20 mgs/kg, i.p.)

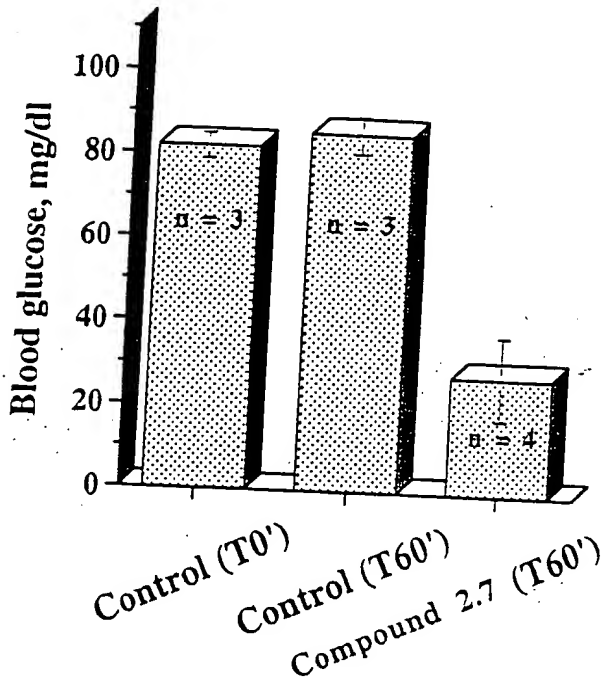

FIGURE 7

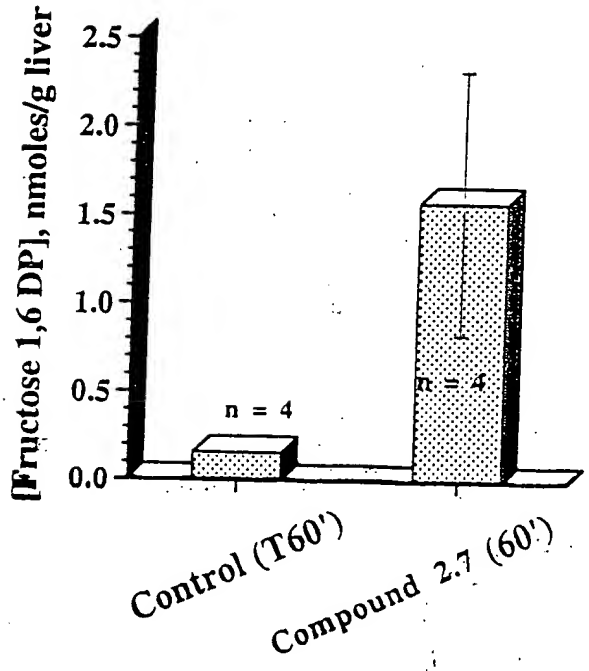

FIGURE 8

24h fasted ZDF Rats + COMPOUND 2.7

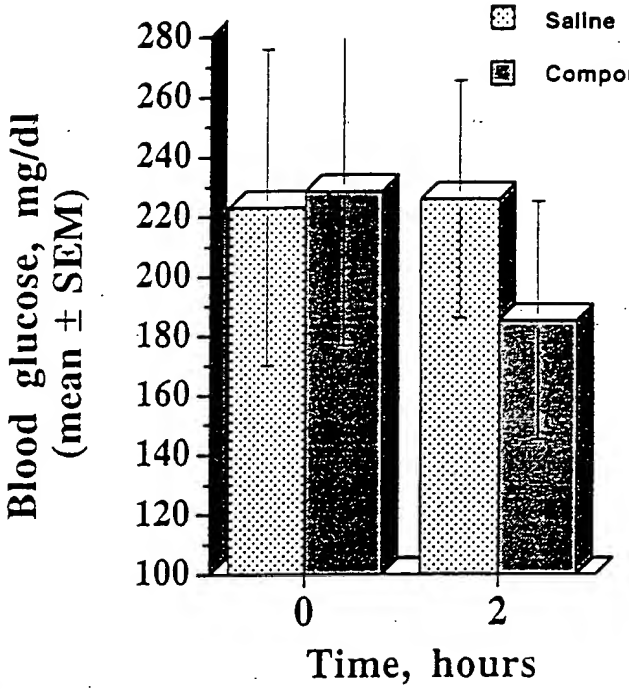

FIGURE 9A

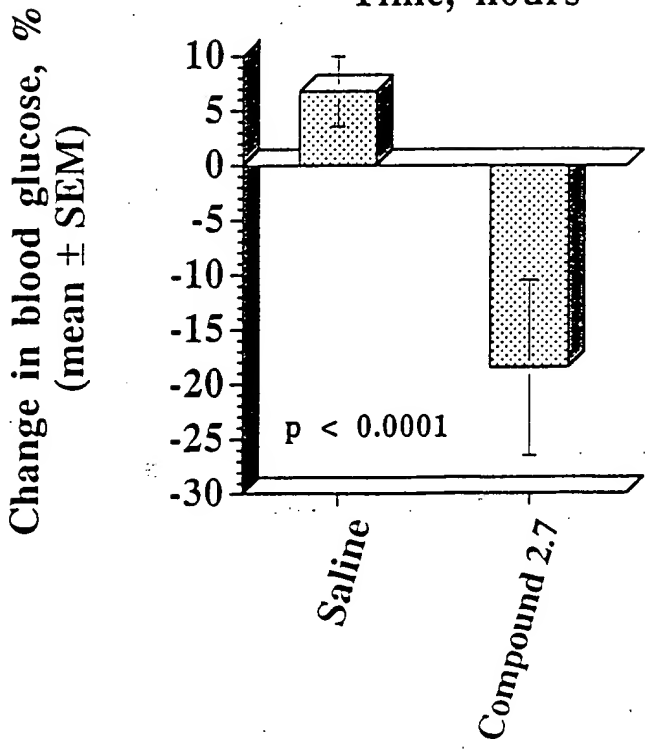

FIGURE 9B

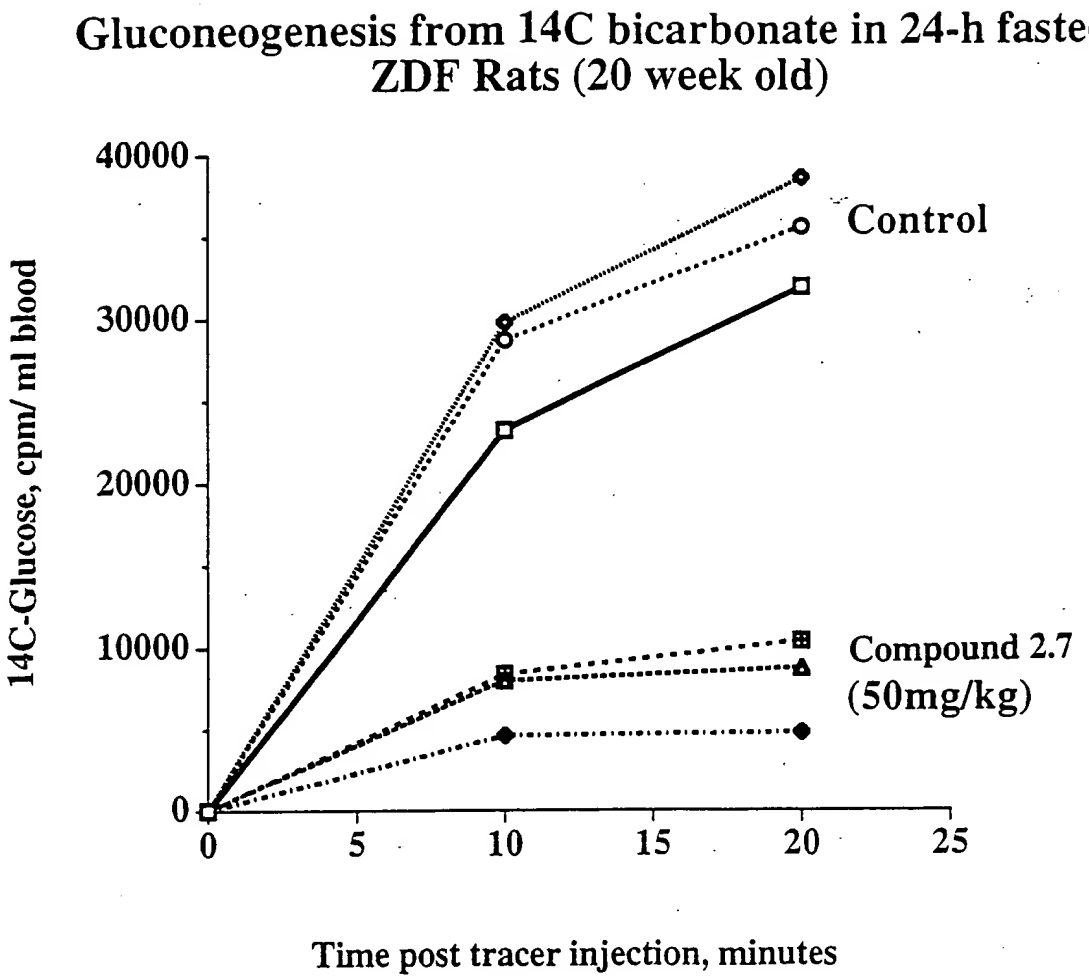
Gluconeogenesis from 14C bicarbonate in 24-h fasted ZDF Rats (20 week old)
14C-Glucose, cpm/ ml blood
40000
30000
20000
10000
0
0
5
10
15
20
25
Time post tracer injection, minutes
Control
Compound 2.7 (50mg/kg)

FIGURE 10

Rat Hepatocytes: Inhibition of Glucose Production and Cellular Penetration

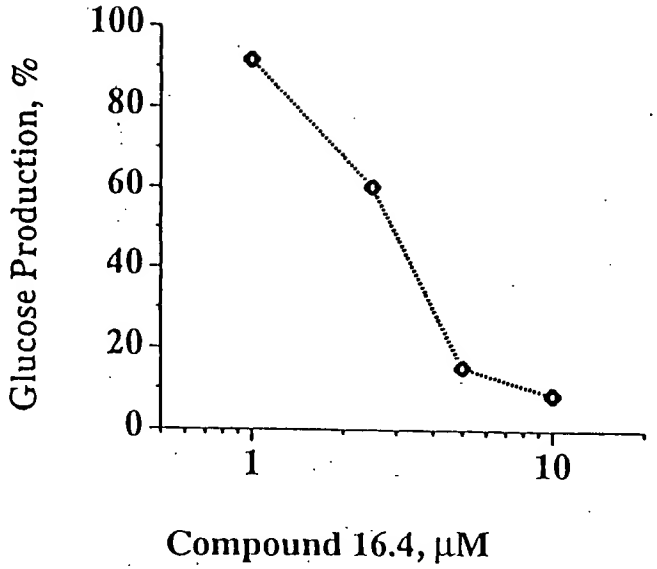

FIGURE 11B

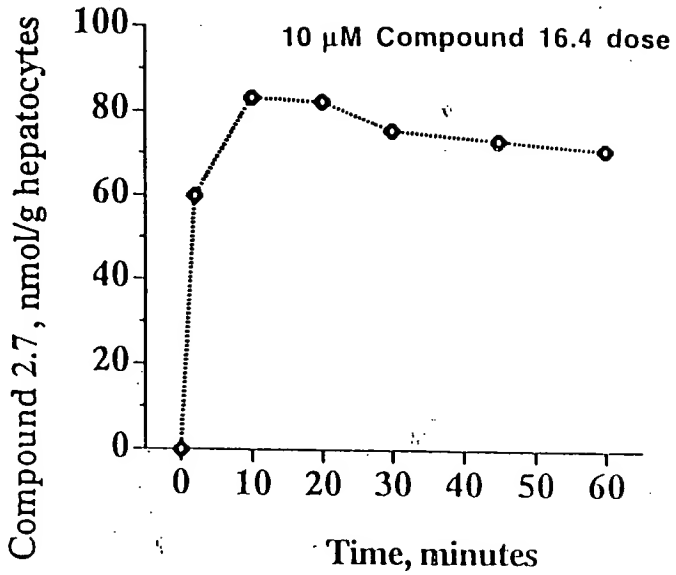

FIGURE 11A